Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

## **CLAIMS**

1. (currently amended) A compound of Formula I:

$$R^{1}-Y-X-N-C-[CH]_{n}-H-N-C-C-N-Ar$$
 $R^{3}$ 
 $O$ 
 $R^{5}$ 
 $R^{7}$ 
 $I$ 

or a pharmaceutically acceptable salt thereof, wherein:

 $R^{1}$  is  $C_6$ - $C_{10}$  aryl substituted with 0-3  $R^{1a}$ , or a  $C_3$ - $C_8$  cycloalkyl substituted with 0-2  $R^{1b}$ , wherein said  $C_3$ - $C_8$  cycloalkyl is saturated or unsaturated;

each  $R^{1a}$  is independently a member selected from the group consisting of H,  $C_1$ - $C_3$  perfluoroalkyl,  $C_3$ - $C_7$  cycloalkyl, F, Cl, Br, CN, NO<sub>2</sub>, OR<sup>10</sup>, SCH<sub>3</sub>, S(=O)CH<sub>3</sub>, S(=O)<sub>2</sub>R<sup>10</sup>, NR<sup>11</sup>R<sup>12</sup>, acetyl, C(=O)OR<sup>13</sup>, C(=O)NR<sup>13</sup>R<sup>14</sup>, S(=O)<sub>2</sub>NR<sup>13</sup>R<sup>14</sup>, phenyl substituted with 0-3 R<sup>15</sup>, and a  $C_1$ - $C_4$  alkyl substituted with 0-2 R<sup>16</sup>;

each  $R^{1b}$  is independently a member selected from the group consisting of H, OH, F, Cl, acetyl, =O,  $C_1$ - $C_6$  alkyl,  $C_1$ - $C_6$  alkoxy,  $CF_3$  and  $OCF_3$ ;

 $R^2$  is a member selected from the group consisting of a phenyl substituted with 0-3  $R^{15}$ , a  $C_1$ - $C_6$  alkyl substituted with 0-2  $R^{2a}$ , wherein said  $C_4$ - $C_6$ -alkyl optionally contains a heteroatom selected from the group consisting of O, S, and  $S(=O)_2$ , a  $C_2$ - $C_6$  alkenyl, a  $C_2$ - $C_6$  alkynyl, a  $C_3$ - $C_7$  cycloalkyl substituted with 0-2  $R^{19}$ , and a  $C_7$ - $C_{11}$  bicycloalkyl substituted with 0-2  $R^{19}$ ;

each  $R^{2a}$  is independently a member selected from the group consisting of a  $C_6$ - $C_{10}$  aryl substituted with 0-3  $R^{15}$ , a  $C_3$ - $C_8$  cycloalkyl substituted with 0-2  $R^{19}$ , and a  $C_7$ - $C_{11}$  bicycloalkyl substituted with 0-2  $R^{19}$ ;

 $R^3$  is a member selected from the group consisting of H and  $C_1$ - $C_4$  alkyl; subscript n is 0 or 1;

 $R^4$  is a member selected from the group consisting of H and  $C_1\text{-}C_6$  alkyl;

 $R^5$  is a member selected from the group consisting of H,  $C_3$ - $C_7$  cycloalkyl,  $C_2$ - $C_6$  alkenyl,  $C_2$ - $C_6$  alkyne, phenyl substituted with 0-2  $R^{15}$ ; and a  $C_1$ - $C_6$  alkyl substituted with 0-2  $R^{18}$ , wherein said  $C_1$ - $C_6$  alkyl optionally contains a heteroatom selected from the group consisting of O, S,  $S(=O)_2$  and  $NR^{17}$ -;

Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

Y is a member independently selected from the group consisting of a bond and  $-(CR^{20}R^{21})_m$ -W- $(CR^{22}R^{23})_p$ -;

subscript p is 1 or 2;

subscript m is 0 or 1;

W is a member-independently selected from the group consisting of a bond, O, S, S(=O),  $S(=O)_2$  and  $NR^{+2}$ -;

X is selected from the group consisting of -C(=O), OC(=O),  $NR^{24}C(=O)$  and  $-S(=O)_2$ ;

each of  $R^6$ ,  $R^7$ ,  $R^8$  and  $R^9$  is independently a member selected from the group consisting of H and  $C_1$ - $C_4$  alkyl;

Ar is a member selected from the group consisting of phenyl substituted with 0-3  $R^{29}$ , and 5 to 6 membered heteroaryl containing 1 to 4 heteroatoms each independently a member selected from the group consisting of N, O and S, wherein said heteroaryl is substituted with 0-3  $R^{29}$ :

each  $R^{10}$  is independently a member selected from the group consisting of H,  $C_3$ - $C_7$  cycloalkyl, a  $C_1$ - $C_3$  perfluoroalkyl, a  $C_1$ - $C_4$  alkyl substituted with 0-1  $R^{25}$ , and a phenyl substituted with 0-3  $R^{15}$ ;

each  $R^{11}$  is independently a member selected from the group consisting of H, 'BOC, Cbz, C<sub>3</sub>-C<sub>8</sub> cycloalkyl, (C<sub>1</sub>-C<sub>6</sub> alkyl)-C(=O)-, (C<sub>1</sub>-C<sub>6</sub> alkyl)-S(=O)<sub>2</sub>- and a C<sub>1</sub>-C<sub>6</sub> alkyl;

each of  $R^{12}$ ,  $R^{13}$  and  $R^{14}$  is independently a member selected from the group consisting of H and  $C_1$ - $C_4$  alkyl;

each  $R^{15}$  is independently a member selected from the group consisting of H, OH, F, Cl, Br, I, CN, NO<sub>2</sub>, COOR<sup>13</sup>, C(=O)NR<sup>13</sup>R<sup>14</sup>, S(=O)<sub>2</sub>NR<sup>13</sup>R<sup>14</sup>, acetyl, -SCH<sub>3</sub>, -S(=O)CH<sub>3</sub>, -S(=O)<sub>2</sub>CH<sub>3</sub>, NR<sup>26</sup>R<sup>27</sup>, C<sub>1</sub>-C<sub>6</sub> alkoxy, C<sub>1</sub>-C<sub>3</sub> perfluoroalkyl, C<sub>1</sub>-C<sub>3</sub> perfluoroalkoxy and a C<sub>1</sub>-C<sub>6</sub> alkyl;

each  $R^{16}$  is independently a member selected from the group consisting of H, OH, COOR<sup>13</sup>, C(=O)NR<sup>13</sup>R<sup>14</sup>, S(=O)<sub>2</sub>NR<sup>13</sup>R<sup>14</sup>, acetyl, -SCH<sub>3</sub>, -S(=O)CH<sub>3</sub>, -S(=O)<sub>2</sub>CH<sub>3</sub>, C<sub>1</sub>-C<sub>6</sub> alkoxy, NR<sup>26</sup>R<sup>27</sup>, and a phenyl substituted with 0-3 R<sup>15</sup>;

R<sup>47</sup> is a member selected from the group consisting of H and C<sub>4</sub>-C<sub>4</sub> alkyl; each R<sup>18</sup> is independently a member selected from the group consisting of H, OH, F, Cl, CN, NO<sub>2</sub>, C(=O)OR<sup>30</sup>, C(=O)NR<sup>13</sup>R<sup>14</sup>, NR<sup>11</sup>R<sup>12</sup>, a C<sub>1</sub>-C<sub>3</sub> perfluoroalkyl, a C<sub>1</sub>-C<sub>3</sub> perfluoroalkoxy, a phenyl substituted with 0-3 R<sup>15</sup>; and C<sub>3</sub>-C<sub>8</sub> cycloalkyl;

Inventors: Liu, et al.

Filing Date: March 23, 2004

Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

each  $R^{19}$  is [[a]] independently a member selected from the group consisting of  $C_1$ - $C_4$  alkyl, F, Cl and  $C_1$ - $C_4$  alkoxy,  $CF_3$  and  $OCF_3$ ;

each of  $R^{20}$ ,  $R^{21}$ ,  $R^{22}$  and  $R^{23}$  is independently a member selected from the group consisting of a bond, H, F, OH,  $C_1$ - $C_4$  alkyl, and  $C_1$ - $C_3$  alkylhydroxy;

R<sup>24</sup> is a member selected from the group consisting of H and C<sub>1</sub> C<sub>4</sub> alkyl;

each  $R^{25}$  is independently a member selected from the group consisting of H,  $C_3$ - $C_7$  cycloalkyl, and a phenyl substituted with 0-3  $R^{15}$ ;

each  $R^{26}$  is independently a member selected from the group consisting of H,  $C_1$ - $C_4$  alkyl,  $(C_1$ - $C_4$  alkyl)-C(=0)- and  $(C_1$ - $C_4$  alkyl)- $S(=0)_2$ -;

each  $R^{27}$  is independently a member selected from the group consisting of H and  $C_1$ - $C_4$  alkyl;

each  $R^{28}$  is independently a member selected from the group consisting of H, a  $C_1$ - $C_6$  alkyl,  $C_3$ - $C_8$  cycloalkyl, a phenyl substituted with 0-3  $R^{15}$ , and a benzyl substituted with 0-2  $R^{15}$ ;

each  $R^{29}$  is independently a member selected from the group consisting of H, F, Cl, Br, I, CN, NO<sub>2</sub>, OR<sup>28</sup>, SR<sup>28</sup>, S(=O)R<sup>28</sup>, S(=O)<sub>2</sub>R<sup>28</sup>, S(=O)<sub>2</sub>NR<sup>13</sup>R<sup>14</sup>, NR<sup>26</sup>R<sup>27</sup>, acetyl, C(=O)NR<sup>13</sup>R<sup>14</sup>, C(=O)OR<sup>13</sup>, C<sub>1</sub>-C<sub>6</sub> alkyl, OCHF<sub>2</sub>, SCF<sub>3</sub>, OCF<sub>3</sub>, and -C(=NH)NH<sub>2</sub>;

alternatively, wherein  $R^{29}$  and  $R^9$  are taken together to form a 5- to 7-membered fused heterocyclic ring containing 1-2 heteroatom(s) each independently a member selected from the group consisting of N, O and S; wherein said 5 to 7 membered fused heterocyclic ring is substituted with 0-2  $R^{19}$ ;

each  $R^{30}$  is independently a member selected from the group consisting of H,  $C_3$ - $C_7$  cycloalkyl,  $C_1$ - $C_4$  alkyl substituted with 0-1  $R^{25}$ , and a phenyl substituted with 0-3  $R^{15}$ ; and with the proviso that  $R^3$ ,  $R^4$ ,  $R^5$ ,  $R^6$ ,  $R^7$ ,  $R^8$ , and  $R^9$  are not all hydrogen.

## 2-3. (canceled)

4. (previously presented) The compound of claim 1, wherein  $R^1$  is phenyl substituted with 0-3  $R^{1a}$ .

## 5-6. (canceled)

Inventors: Liu, et al.

Filing Date: March 23, 2004 Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

7. (currently amended) The compound of claim 9 1, according to formula Ia:

wherein:

 $R^1$  is  $C_3$ - $C_8$  cycloalkyl substituted with 0-2  $R^{1b}$ , wherein said  $C_3$ - $C_8$  cycloalkyl is saturated or unsaturated; and

 $R^2$  is a member selected from the group consisting of a phenyl substituted with 0-3  $R^{15}$ , a  $C_1$ - $C_6$  alkyl substituted with 0-2  $R^{2a}$ , and a  $C_3$ - $C_7$  cycloalkyl substituted with 0-2  $R^{19}$ ; and

Ar is phenyl substituted with 0-3 R<sup>29</sup>, or alternatively, R<sup>29</sup> and R<sup>9</sup> are taken together to form a 5- to 7-membered fused heterocyclic ring containing 1-2 heteroatom(s) each independently a member selected from the group consisting of N, O and S; wherein said 5- to 7-membered fused heterocyclic ring is substituted with 0-2 R<sup>19</sup>.

8. (currently amended) The compound of claim [[7]] 9, wherein:

 $R^2$  is a member selected from the group consisting of a  $C_1$ - $C_2$  alkyl substituted with 1  $R^{2a}$ , and  $C_1$ - $C_6$  alkyl;

each  $R^{2a}$  is independently a member selected from the group consisting of a phenyl substituted with 0-3  $R^{15}$ , and a  $C_3$ - $C_8$  cycloalkyl substituted with 0-2  $R^{19}$ ;

 $R^5$  is a member selected from the group consisting of H,  $C_3$ - $C_7$  cycloalkyl; <u>and</u> a  $C_1$ - $C_6$  alkyl substituted with 0-1  $R^{18}$ , wherein said  $C_1$ - $C_6$ -alkyl optionally contains a heteroatom selected from the group consisting of O, S,  $S(=O)_2$  and  $NR^{17}$ -; and

each  $R^{18}$  is independently a member selected from the group consisting of H, OH, F, Cl, CN, C(=O)OR<sup>30</sup>, C(=O)NR<sup>13</sup>R<sup>14</sup>, NR<sup>11</sup>R<sup>12</sup>, a phenyl substituted with 0-3 R<sup>15</sup>, and C<sub>3</sub>-C<sub>8</sub> cycloalkyl.

9. (currently amended) The compound of claim [[7]] 1, wherein said compound is of the formula:

Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

10-15. (canceled)

16. (currently amended) The compound of claim 18 1, according to formula le

wherein:

 $R^1$  is  $C_6$ - $C_{10}$  aryl substituted with 0-3  $R^{1a}$ ; and

each  $R^{1a}$  is independently a member selected from the group consisting of H,  $C_1$ - $C_3$  perfluoroalkyl,  $C_3$ - $C_7$  cycloalkyl, F, Cl, Br, CN, NO<sub>2</sub>, OR<sup>10</sup>, SCH<sub>3</sub>, S(=O)CH<sub>3</sub>, S(=O)<sub>2</sub>R<sup>10</sup>, NR<sup>11</sup>R<sup>12</sup>, acetyl, C(=O)OR<sup>13</sup>, C(=O)NR<sup>13</sup>R<sup>14</sup>, S(=O)<sub>2</sub>NR<sup>13</sup>R<sup>14</sup>, phenyl substituted with 0-3 R<sup>15</sup>, and a  $C_1$ - $C_4$  alkyl substituted with 0-2 R<sup>16</sup>; and

Ar is phenyl substituted with 0 3 R<sup>29</sup>, or alternatively, R<sup>29</sup> and R<sup>9</sup> are taken together to form a 5 to 7 membered fused heterocyclic ring containing 1-2 heteroatom(s) each independently a member selected from the group consisting of N, O and S; wherein said 5 to 7 membered fused heterocyclic ring is substituted with 0 2 R<sup>19</sup>.

17. (currently amended) The compound of claim [[16]] 18, wherein:

 $R^2$  is a member selected from the group consisting of a  $C_1$ - $C_2$  alkyl substituted with 1  $R^{2a}$ , and  $C_1$ - $C_6$  alkyl;

each  $R^{2a}$  is independently a member selected from the group consisting of a phenyl substituted with 0-3  $R^{15}$ , and a  $C_3$ - $C_8$  cycloalkyl substituted with 0-2  $R^{19}$ ; and

 $R^5$  is a member selected from the group consisting of H,  $C_3$ - $C_7$  cycloalkyl; and a  $C_1$ - $C_6$  alkyl, wherein said  $C_1$ - $C_6$ -alkyl optionally contains a heteroatom selected from the group consisting of O, S,  $S(=O)_2$  and  $NR^{17}$ .

18. (currently amended) The compound of claim [[16]] 1, wherein said compound is of the formula:

Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

19. (currently amended) The compound of claim 9 1, according to formula Ia

wherein:

 $R^{1}$  is  $C_{6}$ - $C_{10}$  aryl substituted with 0-3  $R^{1a}$ ;

each  $R^{1a}$  is independently a member selected from the group consisting of H,  $C_1$ - $C_3$  perfluoroalkyl,  $C_3$ - $C_7$  cycloalkyl, F, Cl, Br, CN, NO<sub>2</sub>, OR<sup>10</sup>, SCH<sub>3</sub>, S(=O)CH<sub>3</sub>, S(=O)<sub>2</sub>R<sup>10</sup>, NR<sup>11</sup>R<sup>12</sup>, acetyl, C(=O)OR<sup>13</sup>, C(=O)NR<sup>13</sup>R<sup>14</sup>, S(=O)<sub>2</sub>NR<sup>13</sup>R<sup>14</sup>, phenyl substituted with 0-3 R<sup>15</sup>; and a  $C_1$ - $C_4$  alkyl substituted with 0-2 R<sup>16</sup>;

 $R^2$  is a member selected from the group consisting of a phenyl substituted with 0-3  $R^{15}$ ; a  $C_1$ - $C_2$  alkyl substituted [[with1 $R^{2a}$ ]] with 1  $R^{2a}$ , and a  $C_3$ - $C_7$  cycloalkyl substituted with 0-2  $R^{19}$ ; and

each  $R^{2a}$  is independently a member selected from the group consisting of a  $C_6$ - $C_{10}$  aryl substituted with 0-3  $R^{15}$ ; a  $C_3$ - $C_8$  cycloalkyl substituted with 0-2  $R^{19}$ ; and a  $C_7$ - $C_{11}$  bicycloalkyl substituted with 0-2  $R^{19}$ ; and

Ar is phenyl substituted with 0-3 R<sup>29</sup>, or alternatively, R<sup>29</sup> and R<sup>9</sup> are taken together to form a 5- to 7-membered fused heterocyclic ring containing 1-2 heteroatom(s) each independently a member selected from the group consisting of N, O and S; wherein said 5- to 7-membered fused heterocyclic ring is substituted with 0-2 R<sup>19</sup>.

20-22. (canceled)

23. (currently amended) The compound of claim 1, according to formula Ig:

$$R^{1}-Y-X-N-C-H-N-C-C-N-N-C-C-N-N-1$$

Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

wherein:

 $R^5$  is a member selected from the group consisting of H,  $C_3$ - $C_7$  cycloalkyl,  $C_2$ - $C_6$  alkenyl,  $C_2$ - $C_6$  alkyne, phenyl substituted with 0-2  $R^{15}$ ; and a  $C_1$ - $C_6$  alkyl substituted with 0-2  $R^{18}$ , wherein said  $C_1$ - $C_6$  alkyl optionally contains a heteroatom selected from the group consisting of O, S,  $S(=O)_2$  and  $NR^{47}$ .

24. (previously presented) The compound of claim 23, according to formula Ih:

$$R^{1}-Y-X-N-C \xrightarrow{H} \stackrel{R^{2}}{-} \stackrel{H}{N} \stackrel{H}{-} \stackrel{H}{C} \stackrel{H}{-} \stackrel{H}{N} \stackrel{H}{-} \stackrel{H}{N} \stackrel{H}{-} \stackrel{H}{N} \stackrel{1}{N} \stackrel{1}{$$

- 25. (currently amended) The compound of claim 1, wherein R<sup>9</sup> is H; and Ar is phenyl substituted with 0-3 R<sup>29</sup>, or alternatively, R<sup>29</sup> and R<sup>9</sup> are taken together to form a 5- to 7-membered fused heterocyclic ring containing 1-2 heteroatom(s) each independently a member selected from the group consisting of N, O and S; wherein said 5- to 7-membered fused heterocyclic ring is substituted with 0-2 R<sup>19</sup>.
  - 26. (canceled)
- 27. (currently amended) A pharmaceutical composition comprising the compound of Formula I in claim 1[[: or]] <u>and</u> a pharmaceutically acceptable <del>salt and an</del> excipient.
- 28. (currently amended) A pharmaceutical composition comprising the compound of claim 38 and a pharmaceutically acceptable excipient.

29-37. (canceled)

- 38. (currently amended) The compound of claim 1, selected from the group consisting of:
- $(S)-N-\{1-[2-(5-Fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-3-methyl-butyl\}-3-methyl-benzamide;$

Inventors: Liu, et al. Filing Date: March 23, 2004

Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

- $N-(S)-\{2-cyclohexyl-1-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl\}-4-phenoxy-benzamide;$
- (S)-3-Cyclohexyl-N-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethyl]-2-[2-(4-methoxy-phenyl)-acetylamino]-propionamide;
- $(S)-N-\{1-[2-(5-Chloro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-3-methyl-butyl\}-3-methyl-benzamide;$
- (S)-N-{3-Cyclohexyl-1-[2-(7-methoxy-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-propyl}-3-methoxy-benzamide;
- $(S)-N-\{3-Cyclohexyl-1-[2-(6-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-propyl\}-3-methoxy-benzamide;$
- (S)-N-{3-Cyclohexyl-1-[2-(7-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-propyl}-3-methoxy-benzamide;
- (S)-N-{3-Cyclohexyl-1-[2-(5-cyano-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-propyl}-3-methoxy-benzamide;
- Cyclopropanecarboxylic acid (S)-{2-cyclohexyl-1-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl}-amide;
- $(S)-N-\{3-Cyclohexyl-1-[2-(4-methoxy-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-propyl\}-3-methoxy-benzamide;$
- (S)-N-{3-Cyclohexyl-1-[2-(5-methoxy-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-propyl}-3-methoxy-benzamide;
- (S)-N-{3-Cyclohexyl-1-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-propyl}-3-methoxy-benzamide;
- (S)-N-{3-Cyclohexyl-1-[2-(5-benzyloxy-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-propyl}-3-methoxy-benzamide;
- N {1 (S) [2 (4 Methoxy phenylamino) propylcarbamoyl] 3 methyl butyl} 3 methyl benzamide:
- N {1 (S) {2 (4 Methoxy phenylamino) 1 methyl ethylcarbamoyl} 3 methyl butyl} 3 methyl benzamide;
- N {1 (S) [2 (4 Methoxy phenylamino) 1 (S) methyl ethylcarbamoyl] 3 methyl butyl} 3 methyl benzamide;
- N {1 (S) [2 (4 Methoxy phenylamino) 1 (R) methyl ethylcarbamoyl] 3 methyl butyl} 3 methyl benzamide;

Application No.: 10/807,613

Inventors: Liu, et al.

PATENT
P1095US10

Inventors: Liu, et al. Filing Date: March 23, 2004

Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

N {2 Cyclohexyl (1S) [2 (4 methoxy phenylamino) (1R) methyl ethylcarbamoyl] ethyl} 3 methoxy benzamide;

 $N-\{(1S)-[2-(5-Fluoro-2,3-dihydro-indol-1-yl)-1,1-dimethyl-ethylcarbamoyl]-2-phenyl-ethyl\}-3-methyl-benzamide;$ 

N {1 (S) [1 (R) Benzyloxymethyl 2 (4 methoxy phenylamino) ethylcarbamoyl] 3-methyl butyl} 3 methyl benzamide;

N (S) {{1 (R) Benzyloxymethyl 2 (5 fluoro 2,3 dihydro indol 1 yl) ethylcarbamoyl} phenyl methyl} 3 methoxy benzamide;

N [1 (S) [1 (R) Benzyloxymethyl 2 (5 fluoro 2,3 dihydro indol 1 yl) ethylcarbamoyl] 2 (4 fluoro phenyl) ethyl] 3 methoxy benzamide;

N {1 (S) [(2 Benzyloxy 1 (R) (5 fluoro 2,3 dihydro indol 1 ylmethyl) ethylcarbamoyl} 3 cyclohexyl propyl} 3 methoxy benzamide;

N-{3-Cyclohexyl-1-(S)-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-1-(R)-hydroxymethylethylcarbamoyl]-propyl}-3-methoxy-benzamide;

N-{3-Cyclohexyl-1-(R)-[(S)-2-(5-fluoro-2,3-dihydro-indol-1-yl)-1-(R)-hydroxymethyl-ethylcarbamoyl]-propyl}-3-methoxy-benzamide;

(S,S)-5-(5-Fluoro-2,3-dihydro-indol-1-yl)-4-[4-methyl-2-(3-methyl-benzoylamino)-pentanoylamino]-pentanoic acid benzyl ester;

(S,S)-5-(5-Fluoro-2,3-dihydro-indol-1-yl)-4-[4-methyl-2-(3-methyl-benzoylamino)-pentanoylamino]-pentanoic acid;

(S,S)-*N*-{1-[3-Carbamoyl-1-(5-fluoro-2,3-dihydro-indol-1-ylmethyl)-propylcarbamoyl]-3-methyl-butyl}-3-methyl-benzamide;

(S,S) N {1 [1 (5 Fluoro 2,3 dihydro indol 1 ylmethyl) 3 ureido propylcarbamoyl] 3 methyl butyl} 3 methyl benzamide;

(S,S)-3-[4-Cyclohexyl-2-(3-methoxy-benzoylamino)-butyrylamino]-4-(5-fluoro-2,3-dihydro-indol-1-yl)-butyric acid benzyl ester;

(S.S)-3-[4-Cyclohexyl-2-(3-methoxy-benzoylamino)-butyrylamino]-4-(5-fluoro-2,3-dihydro-indol-1-yl)-butyric acid;

 $(S,S)-N-\{1-[1-Benzyl-2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-3-cyclohexyl-propyl\}-3-methoxy-benzamide;$ 

(S,S)-N-{3-Cyclohexyl-1-[1-(5-fluoro-2,3-dihydro-indol-1-ylmethyl)-3-methyl-butylcarbamoyl]-propyl}-3-methoxy-benzamide;

Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

- (S,S)-*N*-{3-Cyclohexyl-1-[1-(5-fluoro-2,3-dihydro-indol-1-ylmethyl)-2-methyl-propylcarbamoyl]-propyl}-3-methoxy-benzamide;
- $(S,S)-N-\{3-Cyclohexyl-1-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-1-phenyl-ethylcarbamoyl]-propyl\}-3-methoxy-benzamide;$
- N {1 (S) [2 (R) Benzyloxy 1 (R) (5 fluoro 2,3 dihydro indol 1 ylmethyl) propylcarbamoyl] 3 cyclohexyl propyl} 3 methoxy benzamide;
- N {1 (R) [1 (R) Benzylsulfanylmethyl 2 (5 fluoro 2,3 dihydro indol 1 yl) ethylcarbamoyl] 3 cyclohexyl propyl} 3 methoxy benzamide;
- (S,S) [5 [4 Cyclohexyl 2 (3 methoxy benzoylamino) butyrylamino] 6 (5 fluoro 2,3 dihydro indol 1 yl) hexyl] carbamic acid benzyl ester;
- $N-\{2-cyclohexyl-1-(S)-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl\}-2-(2-fluoro-biphenyl-4-yl)-propionamide;$
- $N-\{2-cyclohexyl-1-(S)-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl\}-ethyl\}-2-p-tolyl-propionamide;$
- $N-\{2-cyclohexyl-1-(S)-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl\}-2-o-tolyl-propionamide;$
- $N-\{2-cyclohexyl-1-(S)-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl\}-2-(4-fluoro-phenyl)-propionamide;$
- 2-(4-Chloro-phenyl)-N-{2-cyclohexyl-1-(S)-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl}-propionamide;
- $N-\{2-cyclohexyl-1-(S)-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl\}-2-(R)-phenyl-propionamide;$
- N-(S)-{2-cyclohexyl-1-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl}-3-methyl-benzamide;
- N-(S)-{2-cyclohexyl-1-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl}-4-(methanesulfonylamino-methyl)-benzamide;
- $N-(S)-\{2-cyclohexyl-1-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl\}-3-methanesulfonyl-benzamide;$
- $N-(S)-\{2-cyclohexyl-1-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl\}-4-methanesulfonylamino-benzamide;$
- $N-\{2-cyclohexyl-1-(S)-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl\}-2-(4-hydroxy-phenyl)-propionamide;$

Application No.: 10/807,613

Inventors: Liu, et al.

PATENT
P1095US10

Filing Date: March 23, 2004

Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

4-Cyclohexyl-N-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethyl]-2-(S)-(2-(R)-phenyl-propionylamino)-butyramide;

 $N-\{2-cyclohexyl-1-(S)-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl]-ethyl\}-2-(R)-phenyl-butyramide;$ 

N {1 (S) [1 (R) Benzyloxymethyl 2 (5 fluoro 2,3 dihydro indol 1 yl) ethylcarbamoyl] 2 cyclohexyl ethyl) 3 methoxy benzamide;

 $N-\{2-Cyclohexyl-1-(S)-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-1-(R)-hydroxymethyl-ethylcarbamoyl]-ethyl\}-3-methoxy-benzamide;$ 

N {1 (S) [1 (R) Benzyloxymethyl 2 (5 fluoro 2,3 dihydro indol 1 yl) ethylcarbamoyl] 3,3 dimethyl butyl} 3 methoxy benzamide;

 $N-\{1-(S)-[2-(5-Fluoro-2,3-dihydro-indol-1-yl)-1-(R)-hydroxymethyl-ethylcarbamoyl\}-3,3-dimethyl-butyl\}-3-methoxy-benzamide;$ 

3 (S) (2 (S) Benzyloxycarbonylamino 4,4 dimethyl pentanoylamino) 4 (5 fluoro 2,3 dihydro indol 1 yl) butyric acid tert butyl ester;

3 (S) (2 (S) Benzyloxycarbonylamino 4,4 dimethyl pentanoylamino) 4 (5 fluoro-2,3 dihydro indol 1 yl) butyric acid;

4-(5-Fluoro-2,3-dihydro-indol-1-yl)-3-(S)-[2-(S)-(3-methoxy-benzoylamino)-4,4-dimethyl-pentanoylamino]-butyric acid tert-butyl ester;

3-(S)-[3-Cyclohexyl-2-(S)-(3-methoxy-benzoylamino)-propionylamino]-4-(5-fluoro-2,3-dihydro-indol-1-yl)-butyric acid benzyl ester;

3-(S)-[3-Cyclohexyl-2-(S)-(3-methoxy-benzoylamino)-propionylamino]-4-(5-fluoro-2,3-dihydro-indol-1-yl)-butyric acid;

4-(5-Fluoro-2,3-dihydro-indol-1-yl)-3-(S)-[2-(S)-(3-methoxy-benzoylamino)-4,4-dimethyl-pentanoylamino]-butyric acid ethyl ester;

 $N-\{1-(S)-[2-Cyano-1-(S)-(5-fluoro-2,3-dihydro-indol-1-ylmethyl)-ethylcarbamoyl]-3,3-dimethyl-butyl\}-3-methoxy-benzamide;$ 

 $N-\{1-(S)-[5-Amino-1-(S)-(5-fluoro-2,3-dihydro-indol-1-ylmethyl)-pentylcarbamoyl]-3-cyclohexyl-propyl\}-3-methoxy-benzamide;$ 

3 (S) (2 (S) Benzyloxycarbonylamino 3 cyclohexyl propionylamino) 4 (5 fluoro 2,3 dihydro indol 1 yl) butyric acid benzyl ester;

1 (S) [1 (R) Benzyloxymethyl 2 (5 fluoro 2,3 dihydro indol 1 yl) ethylcarbamoyl] 2 cyclohexyl ethyl) carbamic acid benzyl ester;

Inventors: Liu, et al.

Filing Date: March 23, 2004

Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

N {3 Cyclohexyl 1 (S) [2 (3,5 dimethoxy benzyloxy) 1 (R) (5 fluoro 2,3 dihydro-indol 1 ylmethyl) ethylcarbamoyl] propyl} 3 methoxy benzamide;

PATENT

4-{2 (R) [4 Cyclohexyl 2 (S) (3 methoxy benzoylamino) butyrylamino] 3 (5 fluoro-2,3 dihydro indol 1 yl) propoxymethyl} benzoic acid methyl ester;

 $(S,S)-N-\{3-Cyclohexyl-1-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-1-(4-hydroxy-benzyl)-ethylcarbamoyl\}-3-methoxy-benzamide;$ 

{2 Cyclohexyl 1 (S) [2 (5 fluoro 2,3 dihydro indol 1 yl) 1 (S) methylethylearbamoyl] ethyl} carbamic acid benzyl ester;

4 Benzyloxy N (R,S) [[2 (4 amidinophenylamino) 1 (S) methyl ethylcarbamoyl] (2,4 dichloro phenyl) methyl} benzamide;

{1 (S) [2 (5 Fluoro 2,3 dihydro indol 1 yl) 1 (S) methyl ethylcarbamoyl] 3,3 dimethyl butyl} carbamic acid benzyl ester;

Cyclopropanecarboxylic acid  $\{1-(S)-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-1-(S)-methyl-ethylcarbamoyl]-3,3-dimethyl-butyl\}-amide;$ 

(S,S) 2 (3 Chloro benzenesulfonylamino) 3 cyclohexyl N [1 methyl 2 (4 trifluoromethoxy phenylamino) ethyl] propionamide;

(S,S) 3 Cyclohexyl N [1 methyl 2 (4 trifluoromethoxy phenylamino) ethyl] 2 (3 trifluoromethoxy benzenesulfonylamino) propionamide;

 $N-((S)-(2-(5-fluoroindolin-1-yl)ethylcarbamoyl)(cyclohexyl)\ methyl)-3-methylbenzamide;$ 

N-((S)-1-(2-(5-fluoroindolin-1-yl)ethylcarbamoyl)-2-(2-chlorophenyl)ethyl)-3-methylbenzamide;

N-((S)-1-(2-(5-fluoroindolin-1-yl)ethylcarbamoyl)-2-(3-chlorophenyl)ethyl)-3-methylbenzamide;

N-((S)-1-(2-(5-fluoroindolin-1-yl)ethylcarbamoyl)-2-(4-chlorophenyl)ethyl)-3-methylbenzamide;

 $(S)-N-\{2-Cyclopentyl-1-[2-(5-fluoro-2,3-dihydro-indol-1-yl)-ethylcarbamoyl\}-ethyl\}-3-methyl-benzamide;$ 

N-((S)-1-(2-(5-fluoroindolin-1-yl)ethylcarbamoyl)-3,3-dimethylbutyl)-3-methylbenzamide;

N-((S)-1-(2-(5-fluoroindolin-1-yl)ethylcarbamoyl)-3-cyclohexylpropyl)-3-methylbenzamide;

PATENT

Application No.: 10/807,613

Inventors: Liu, et al. Filing Date: March 23, 2004

Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

- N-((S)-1-(2-(5-fluoroindolin-1-yl)ethylcarbamoyl)-2-phenylethyl)-3-methylbenzamide;
- N-(R,S)-((3-(5-fluoroindolin-1-yl)-1-hydroxypropan-2-(R)-ylcarbamoyl)(2,4-dichlorophenyl)methyl)-3,4-difluorobenzamide;
- N (S) ((3 (benzyloxy) 1 (5-fluoroindolin 1-yl)propan 2 (R) ylcarbamoyl)(2,4-dichlorophenyl)methyl) 3,4-difluorobenzamide;
- (R,S)-N-((2-(5-fluoroindolin-1-yl)ethylcarbamoyl)(2,4-dichlorophenyl)methyl)-3-methylbenzamide;
- (S,S)-N-((3-(5-fluoroindolin-1-yl)-1-hydroxypropan-2-ylcarbamoyl)(2,4-dichlorophenyl)methyl)-3,4-difluorobenzamide;
- (S,S)-4-(5-Fluoro-2,3-dihydro-indol-1-yl)-3-[2-(3-methoxy-benzoylamino)-4,4-dimethyl-pentanoylamino]-butyric acid; and
- (S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro indol 1 yl) ethyl] 2 (5 isoxazol 3 yl-thiophene 2 sulfonylamino) propionamide;
- (S) 2 (3 Biphenyl 4 yl ureido) 3 cyclohexyl N [2 (5 fluoro 2,3 dihydro indol 1 yl) ethyl] propionamide;
- (S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro mdol 1 yl) ethyl] 2 (4 phenoxybenzenesulfonylamino) propionamide:
- (S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro indol-1 yl) ethyl] 2 (naphthalene 1 sulfonylamino) propionamide;
- (S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro midol 1 yl) ethyl] 2 (4 trifluoromethylbenzenesulfonylamino) propionamide;
- (S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro indol 1 yl) ethyl] 2 (4-trifluoromethoxy benzenesulfonylamino) propionamide:
- (S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro indol 1 yl) ethyl] 2 [4 (4 fluoro phenoxy) benzenesulfonylamino] propionamide;
- (S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro indol 1 yl) ethyl] 2 (4' methoxy-biphenyl 4 sulfonylamino) propionamide:
- (S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro mdol 1 yl) ethyl] 2 (4 methoxy-benzenesulfonylamino) propionamide;
- (S) 3 Cyclohexyl 2 (4 difluoromethoxy benzenesulfonylamino) N [2 (5 fluoro 2.3 , dihydro indol 1 yl) ethyl] propionamide: ;

Inventors: Liu, et al.

Filing Date: March 23, 2004

Response to Final Office Action mailed August 2, 2007

Date: September 24, 2007

- (S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro indol 1 yl) ethyl] 2 phenylmethanesulfonylamino propionamide;
- (S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro indol 1 yl) ethyl] 2 (toluene 3 sulfonylamino) propionamide;
- (S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro indol 1 yl) ethyl] 2 [4 (4 methoxy-phenoxy) benzenesulfonylamino] propionamide;

PATENT

- (S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro indol-1 yl) ethyl] 2 (3 methoxy-benzenesulfonylamino) propionamide;
- (S,S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro indol 1 yl) 1 methyl ethyl] 2 (toluene 3 sulfonylamino) propionamide;
- (S,S) 3 [4,4 Dimethyl 2 (toluene 3 sulfonylamino) pentanoylamino] 4 (5 fluoro 2,3-dihydro indol 1 yl) butyric acid tert butyl ester;
- (S,S) 3 Cyclohexyl N [2 (5 fluoro 2,3 dihydro indol 1 yl) 1 methyl ethyl] 2 (3-trifluoromethoxy benzenesulfonylamino) propionamide;
- (S,S) 2 (3 Chloro benzenesulfonylamino) 3 cyclohexyl N [2 (5 fluoro 2,3 dihydro indol 1 yl) 1 methyl ethyl] propionamide;
- (S,S)-N-{3-Cyclohexyl-1-[1-(5-fluoro-2,3-dihydro-indol-1-ylmethyl)-3-hydroxy-propylcarbamoyl]-propyl}-3-methoxy-benzamide;
- (S,S) 3 [4,4 Dimethyl 2 (toluene 3 sulfonylamino) pentanoylamino] 4 (5 fluoro 2,3 dihydro indol 1 yl) butyric acid;
- (S,S) 2 Benzenesulfonylamino 3 cyclohexyl N [2 (5 fluoro 2,3 dihydro indol 1 yl) 1 methyl ethyl] propionamide; and
- (S,S) 4,4 Dimethyl 2 (toluene 3 sulfonylamino) pentanoic acid [2 (5 fluoro 2,3-dihydro indol 1 yl) 1 methyl ethyl] amide.